

WORLD INTELLECTUAL PROPERTY ORGANIZATION International Bureau



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60/085,925 18 May 1998 (18.05.98) 60/085,928 18 May 1998 (18.05.98) 60/085,920 18 May 1998 (18.05.98) (71) Applicant (for all designated States except US): GENOME SCIENCES, INC. [US/US]; 9410 K Avenue, Rockville, MD 20850 (US). (72) Inventors; and (75) Inventors/Applicants (for US only): RUBEN, States (US/US); 18528 Heritage Hills Drive, Olney, M	U HUMA Key We	3142 Quesada Street, N.W., Washington, DC 20015 (US ENDRESS, Gregory, A. [US/US]; 9729 Clagett Fai Drive, Potomac, MD 20854 (US). EBNER, Reinha [DE/US]; 9906 Shelburne Terrace #316, Gaithersburg, M 20878 (US). (74) Agents: BROOKES, A., Anders et al.; Human Genor Sciences, Inc., 9410 Key West Avenue, Rockville, M 20850 (US).	
(corco), 10320 Heritage This Brive, Onley, in	200.	(81) Designated States: AL, AM, AT, AU, AZ, BA, BB, BG, B BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, G GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, K KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, M MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, S SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZV ARIPO patent (GH, GM, KE, LS, MW, SD, SL, SZ, U ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, T TM), European patent (AT, BE, CH, CY, DE, DK, ES, I FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI pate (BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, N SN, TD, TG).	
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(54) Title: 97 HUMAN SECRETED PROTEINS

(57) Abstract

The present invention relates to novel human secreted proteins and isolated nucleic acids containing the coding regions of the genes encoding such proteins. Also provided are vectors, host cells, antibodies, and recombinant methods for producing human secreted proteins. The invention further relates to diagnostic and therapeutic methods useful for diagnosing and treating disorders related to these novel human secreted proteins.

amendments.

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AAZ65250 to AAZ65350 represent 97 isolated human secreted protein genes. This sequence was found to be present on human chromosome 3.

AAY76124 to AAY76223 represent the secreted proteins encoded by the 97 human genes. The genes and their corresponding secreted polypeptides are useful for preventing, treating or ameliorating medical conditions, e.g. by protein or gene therapy. Also pathological conditions can be diagnosed by determining the amount of the new polypeptides in a sample or by determining the presence of mutations in the new genes. Specific uses are described for each of the 97 genes, based on which tissues they
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Human; secreted protein; cancer; tumour; developmental abnormality; foetal deficiency; blood disorder; immune system disorder; inflammation; autoimmune disease; allergy; Alzheimer's disease; cognitive disorder; schizophrenia; arthritis; asthma; psoriasis; sepsis; skin disorder; atherosclerosis; diabetes; cardiovascular disorder; kidney disorder; disestive disorder; endocrine disorder; infection; AIDS; leukaemia; therapy; chromosome 3; ds.
                                                                                                                                                                               New isolated human genes and the secreted polypeptides they encode, useful for diagnosis and treatment of e.g. cancers, neurological disorders, immune diseases, inflammation or blood disorders
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P-PSDB; AAY76143.
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195 / ACI

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8888888888888888 alignment_block: alignment_scores: US-09-899-471-2 x AAZ65269 Quality: 1671.00 Ratio: 4.157 Percent Similarity: 81.048 Align seg 1/1 to: AAZ65269 777 163 uGlyAlaGluValGlnIleTrpSerTyrThrLysProArgTyrGlnLysG ||||:::||||||:::|||| | 127 AGGGAGTGAGGTACGAATCTGGTCCTATACTCAGCCCAGGTACGAGAAGG 180 527 GATGAGGAAAAGTTTGGAGGAGCAGCTGACTTAGGGGGTGGAGGAGCCTAG 477 427 377 327 are most highly expressed in, and include developing products for the diagnosis or treatment of cancer, tumours, developmental abnormalities and foetal deficiencies, blood disorders, diseases of the immune system, autoimmune diseases, inflammation, allergies, Alzhelmer's and cognitive disorders, schizophrenia, arthritis, asthma, psoriasis, sepsis, skin disorders, schizophrenia, arthritis, asthma, psoriasis, sepsis, skin disorders, atherosclerosis, diabettes, cardiovascular disorders, kidney disorders, digestive/endocrine disorders, infections and AIDS. The polypeptides are also useful for identifying their binding partners. The sequences shown in AAY76224 to AAY76424 represent fragments of the security of the 189 101 Glu......AlaGlyLysSerAspSerGluLeuGlnGluSerAr 113 277 84 alargvalvalvalhisLeualavalhisGlyhisTrpalaGluProGlu 100 ::|||||| || |||||||| 67 34 erLeuGlyLeuSerCysHisLeuTrpAspGlyAspValLeuCysLeuPro 50 17 lValValSerLeuGluArgLeuMetGluProGlnAspThrAlaArgCysS 34 Sequence 1752 BP; 324 A; secreted proteins. TGCGTGTGGMTGTCCACTTGGCCGTGCATGGGCACTGGGAAGAGCCTGAA nThrGluLeuValLeuArgCysProGlnLysThrAspCysAlaLeuArgV luLeuAsnLeuThrGlnGlnLeuPro.... GACAGAGCTGGTGCTGAGGTGCCAGAAGGAGACCGACTGTGACCTCTGTC 476 GGTCCTTTCTCTGGAGAGGCCTTGTGGGGCCTCAGGACGCTACCCACTGCT AAYTCAACCACACAGCAGCTGCCTGACTGCAGGGGGGCTCGAAGTCTGG 555 C; from: 1 Length: 496
Gaps: 5
Percent Identity: 65.726 522 G; 347 T; 4 other; to: 1752 163 826 180 146 576 526 426 376 84 67 188 776 726 676 326 276

As

seq_name:

/SIDS1/gcgdata/geneseq/geneseqn-embl/NA2001A.DAT:AAC85028

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.euLeuGlnAspPheArgSerHisGlnCysMetGlnLeuTrpAsn ::::: TACTACAAGACCTGCAGTCAGGCCAGTGTCTGCAGCTATGG AsnMetGlySerLeuTrpAlaCysProMetAspLysTyrIleHi
9 sThrProLeuProSerMetAlaSerThrArgAlaAlaArgLeuGlyGl
3 LysThrGlyLeuAsnAsnThrSerValCysAlaLeuGluProSerGlyC :;: 6 CGAGGCCCCCAGGACAGATCCCTCTGTGCCTTGGAACCCAGTGGCT
6 rpAlaAspSerLeuGlyProPheLysAspAspMetLeuLeuValGluMe
9 uCysValGlnValSerThrTrpGluLysValGlnLeuGlnAlaCysSer
3 ThrValasnGluProGlnAspPheGlnLeuValalaGlyHisProAsnL :::::
6 sp6lnSerProCysGlnProLeuValProProValProGlnLysAsnAl
9 phlaproCysCysLeuProGlyLysValThrLeuCysTrpGlnAlapro
3 TrpHisileAlaArgLeuArgValLeuSerProGlyValTrpGlnLeuA ::: ::: :::
6 rdvalGluPheCysProPheArgGluAspProGlyAlaHisArgAsnLeu
39 uValProCysLeuCysIleGlnValTrpSerLeuGluProAspSerGluA
23 TyrLysAsnLeuThrGlyProGlnAsnIleThrLeuAsnHisThrAspL :::
06 erPheLeuLeuTyrLeuArgProValProAspAlaLeuLysSerLeuTr ::::: ::: 27 GCTCTCCCTGTACTGGAATCAGGTCCAGGGCCCCCAAAACCCCGGTG
9 pdlyaspasnvalleuLeuThrLeuAspValSerGluGluGlnaspPhe
27 AACAGCATCCCGAGCTGCTGGGCCCTGGCCTCAACGTGTCAGCAG

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chromosome 3; ds.
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(HUMA-) HUMAN GENOME SCI INC.

Olsen SM, DW, Shi Y, Florence Endress GA, Young PE, Weiss GA, Ebner R; ,× N1 J, Rosen CA, Carter g PE, Wei F, Brewer LA, Soppet Moore PA;

2000-062296/05.

New isolated human genes and the secreted polypeptides they encode, useful for diagnosis and treatment of e.g. cancers, neurological disorders, immune diseases, inflammation or blood disorders -

Claim 1; Page 308; 475pp; English.

This sequence was found to be present on human chromosome 3.

C AN776124 to AN776223 represent the secreted proteins encoded by the 97 change genes. The genes and their corresponding secreted polypeptides are useful for preventing, treating or amellorating medical conditions, be e.g. by protein or gene therapy. Also pathological conditions can be considered by determining the amount of the new polypeptides in a sample core by determining the presence of mutations in the new genes. Specific cuses are described for each of the 97 genes, based on which tissues they care most highly expressed in, and include developing products for the candidations of the instance of the instance of the instance of the instance system, and foetal deficiencies, blood disorders, diseases of the immune system, cautoimmune diseases, inflammation, allergies, Alzheimer's and cognitive constructors, atherosclerosis, diabetes, cardiovascular disorders, kidney collopeptides are also useful for identifying their binding partners. The sequences shown in AAY76224 to AAY76424 represent fragments of the to AAZ65350 represent 97 isolated human secreted protein genes

Sequence 1752 BP; 324 A; 555 ü 522 <u>.</u> 347 T; 4 other;

alignment_scores:

Percent Similarity: Quality: Ratio: : 1801.00 : 4.248 85.484 Percent Identity: Length: Gaps: 69 . 556

alignment_block: US-09-899-471-5 x AAZ65269

Align seg 1/1 to: AAZ65269 from: 1 to: 1752

- 17 1 MetProValSerTrpPheLeuLeuSerLeuAlaLeuGlyArgAsnProVa ATGCCTGTGCCCTGGTTCTTGCTGTCCTTGGCACTGGGCCGAAGCCCAGT 276 17
- lValValSerLeuGluArgLeuMetGluProGlnAspThrAlaArgCysS GGTCCTTTCTCGGAGAGGCCTTGTGGGGCCTCAGGACGCTACCCACTGCT 34
- 50
- 327 51 GlySerLeuGlnSerAlaProGlyProValLeuValProThrArgLeuGl 67 376
- 377 GGGGACATCGTGCCTGCTCCGGGCCCCCGTGCTGCGCCCTACGCACCTGCA 426
- 427 67 GACAGAGCTGGTGCTGAGGTGCCAGAAGGAGACCGACTGTGACCTCTGTC nThrGluLeuValLeuArgCysProGlnLysThrAspCysAlaLeuArgV 476 84
- alargValValHisLeuAlaValHisGlyHisTrpAlaGluProGlu::||||||| TGCGTGTGGMTGTCCACTTGGCCGTGCATGGGCACTGGGAAGAGCCTGAA
- Glu... .AlaGlyLysSerAspSerGluLeuGlnGluSerAr

1176 1077 1027 1376 1326 1276 1126 :::|||||||||||||||||||||||||||||||||:::
1226 GTGGGGACCCTGCCAGCCACCGGTTTCCTGGGAGAAYGTC 827 577 113 527 977 927 877 197 777 180 727 163 677 147 627 130 397 297 280 263 247 230 213 ProGlyGlnSerValGlySerAlaValPheAspCysPheGluAlaSerLe gAsnAlaSerLeuGlnAlaGlnValValLeuSerPheGlnAlaTyrProI 130 leAlaArgCysAlaLeuLeuGluValGlnValProAlaAspLeuValGln 146 GATGAGGAAAAGTTTGGAGGAGCAGCTGACTTAGGGGGTGGAGGAGCCTAG GGCCTGACTCCCTGGGGCCTCTCAAAGACGATGTGCTACTGTTGGAGACA 1425 CTGTGTTCAGGTGAACAGCTCGGAGAAGCTGCAGCTGCAGGAGTGCTTGT GACG uGl|yAlaGluValGlnIleTrpSerTyrThrLysProArgTyrGlnLysG GAATGCCTCTCCCAGGCCCAAGTCGTGCTCCTCCTTCCAGGCCTACCCTA 626 LysThrGlyLeuAsnAsnThrSerValCysAlaLeuGluProSerGlyCy 413 uCysValGlnValSerThrTrpGluLysValGlnLeuGlnAlaCysSerT spGlnSerProCysGlnProLeuValProProValProGlnLysAsnAla 346 TrpHisIleAlaArgLeuArgValLeuSerProGlyValTrpGlnLeuAs
(||::: ||||||||||:::|||::: AACAGCATCCCGAGCTGCTGGGCCCTGCCCTGGCTCAACGTGTCAGCAGA luLeuAsnLeuThrGlnGlnLeuProAspCysArgGlyLeuGluValArg TTTGGTCAGTCTGTGGCTCTGTGGTATATGACTGCTTCGAGGCTGCCCT CTGCCCGCTGCGTGCTGGAGGTGCAAGTGCCTGCTGCCCCTTGTGCAG rpAlaAspSerLeuGlyProPheLysAspAspMetLeuLeuValGluMet 396 ACTGTGGACAAGGTTCTCGAGTTCCCATTGCTGAAAGGCCACCCTAACCT ThrValAsnGluProGlnAspPheGlnLeuValAlaGlyHisProAsnLe CGCACCGTGCTCGCTGCCCGCAGAAGCGGCACTGTGCTGGCGGGGCTCCGG pAlaProCysCysLeuProGlyLysValThrLeuCysTrpGlnAlaProA 330 rgValGluPheCysProPheArgGluAspProGlyAlaHisArgAsnLeu GGTTCCCTGCCTCTGTATTCAGGTGTGGCCTCTGGAACCTGACTCCGTTA TyrLysAsnLeuThrGlyProGlnAsnIleThrLeuAsnHisThrAspLe GCCTCTCCCTGTACTGGAATCAGGTCCAGGGCCCCCCAAAACCCCCGGTGG erPheLeuLeuTyrLeuArgProValProAspAlaLeuLysSerLeuTrp AspSerIleGlnSerCysTrpValLeuProTrpLeuAsnValSerThrAs AAYTCAACCACACACAGCAGCTGCCTGACTGCAGGGGGGCTCGAAGTCTGG . AACATCTGCCCCTTCAGGNAGGACCCCCGCGCACACCAGAACCTC 363 1275 1225 296 1076 280 1026 263 976 246 926 876 826 196 776 180 726 163 676 576 1125